

EAST Search History

Ref #	Hits	Search Query	DBs	Default Operator	Plurals	Time Stamp
L2	457	436/79.ccls.	US-PGPUB; USPAT; USOCR	OR	ON	2008/01/01 18:33
L3	394	436/79.ccls.	US-PGPUB; USPAT	OR	ON	2008/01/01 18:33
L4	368	436/79.ccls.	USPAT	OR	ON	2008/01/01 18:34
L5	11	bapta and "544".clas.	USPAT	OR	ON	2008/01/01 18:44
L6	22	bapta and "546".clas.	USPAT	OR	ON	2008/01/01 18:44

=> d his

(FILE 'HOME' ENTERED AT 09:56:09 ON 01 JAN 2008)
FILE 'CA' ENTERED AT 09:56:15 ON 01 JAN 2008
L1 6982 S (NEAR OR ADJACENT OR PROXIM?) (3A) MEMBRANE
L2 185032 S (CALCIUM OR CA OR CA2 OR MAGNESIUM OR MG2 OR INDICATOR OR
FLUOROCHRO? OR FLUOROPHOR?) (5A) (DETECT? OR DETERMIN? OR REPORT?
OR OPERAT? OR TEST? OR ANALY? OR ASSAY? OR MEASUR? OR MONITOR? OR
SENSE# OR SENSOR OR SENSING OR PROBE# OR PROBING OR QUANTITAT? OR
QUANTIF? OR QUANTA?)
L3 16 S L1(4A) (INDICATOR OR FLUOROPHOR? OR FLUOROCHRO?)
L4 218 S L1 AND L2
L5 34 S L1 AND (PIPERAZ? OR ZWITTER?)
L6 179 S L4 AND PY<2004
L7 43 S L1(8A) L2
L8 6 S L6 AND FFP?
L9 16 S L4/TI, IT, ST
L10 87 S L3, L5, L7-9
FILE 'BIOSIS' ENTERED AT 10:16:02 ON 01 JAN 2008
L11 82 S L10
FILE 'MEDLINE' ENTERED AT 10:16:40 ON 01 JAN 2008
L12 68 S L10
FILE 'CA, BIOSIS, MEDLINE' ENTERED AT 10:17:19 ON 01 JAN 2008
L13 113 DUP REM L10 L11 L12 (124 DUPLICATES REMOVED)

=> d bib, ab, kwic l13 1-113

L13 ANSWER 35 OF 113 CA COPYRIGHT 2008 ACS on STN
AN 139:347654 CA
TI **Near-membrane** iminocoumarin-based low affinity fluorescent Ca²⁺
indicators
AU Liepouri, F.; Deligeorgiev, T. G.; Veneti, Z.; Savakis, C.;
Katerinopoulos, H. E.
CS Department of Chemistry, University of Crete, Crete, 71 409, Greece
SO Cell Calcium (2002), 31(5), 221-227
AB Two new potential **near-membrane** iminocoumarin-based fluorescent Ca²⁺
indicators were synthesized and the spectral profiles of their free and
Ca²⁺ bound forms were studied. The probes incorporate in their BAPTA-
related structures, the 3-(benzimidazolyl)iminocoumarin or the 3-
(benzothiazolyl)iminocoumarin moiety, substituted at the imino nitrogen
with an n-dodecyl lipophilic chain. The compds. are excited with
visible light and have Ca²⁺ dissocn. const. values of 5.50 and 4.49 μ M,
resp., the highest reported to date in the literature. Fluorescence
spectra studies indicated a clear shift in their excitation wavelength
maxima upon Ca²⁺ binding along with changes in fluorescence intensity
that enable the compds. to be used as ratiometric **near-membrane**, low Ca²⁺
+ affinity **probes**.

L13 ANSWER 66 OF 113 CA COPYRIGHT 2008 ACS on STN
AN 125:52738 CA
TI **Near-membrane** [Ca²⁺] transients resolved using the Ca²⁺ indicator **FFP18**
AU Etter, Elaine F.; Minta, Akwasi; Poenie, Martin; Fay, Fredric S.
CS Dep. Physiology Biomedical Imaging Group, Univ. Massachusetts Med.

Center, orcester, MA, 01605, USA

SO Proceedings of the National Academy of Sciences of the United States of America (1996), 93(11), 5368-5373

AB Ca^{2+} -sensitive processes at cell membranes involved in contraction, secretion, and neurotransmitter release are activated in situ or in vitro by Ca^{2+} concns. [Ca^{2+}] 10-100 times higher than [Ca^{2+}] measured during stimulation in intact cells. This paradox might be explained if the local [Ca^{2+}] at the cell membrane is very different from that in the rest of the cell. Sol. Ca^{2+} indicators, which indicate spatially averaged cytoplasmic [Ca^{2+}], cannot resolve these localized, **near-membrane** [Ca^{2+}] signals. **FFP18**, the newest Ca^{2+} indicator designed to selectively **monitor near-membrane** [Ca^{2+}], has a lower Ca^{2+} affinity and is more water sol. than previously used membrane-assocg. Ca^{2+} indicators. Images of the intracellular distribution of **FFP18** show that >65% is located on or **near** the plasma **membrane**. [Ca^{2+}] transients recorded using **FFP18** during membrane depolarization-induced Ca^{2+} influx show that **near-membrane** [Ca^{2+}] rises faster and reaches micromolar levels at early times when the cytoplasmic [Ca^{2+}], recorded using fura-2, has risen to only a few hundred nanomolar. High-speed series of digital images of [Ca^{2+}] show that **near-membrane** [Ca^{2+}], **reported by FFP18**, rises within 20 ms, peaks at 50-100 ms, and then declines. [Ca^{2+}] **reported** by fura-2 rose slowly and continuously throughout the time images were acquired. The existence of these large, rapid increases in [Ca^{2+}] directly beneath the surface membrane may explain how numerous Ca^{2+} -sensitive membrane processes are activated at times when bulk cytoplasmic [Ca^{2+}] changes are too small to activate them.

L13 ANSWER 71 OF 113 CA COPYRIGHT 2008 ACS on STN

AN 125:7754 CA

TI **Near membrane** Ca^{2+} changes resulting from store release in neutrophils: detection by **FFP-18**

AU Davies, E. V.; Hallett, M. B.

CS Mol. Signalling Group, Univ. Wales, Cardiff, UK

SO Cell Calcium (1996), 19(4), 355-362

AB **FFP-18** was incorporated into the inner face of the plasma membrane of human neutrophils by incubation with its acetoxymethyl ester. Conversion to the Ca^{2+} sensitive intracellular **indicator** was **monitored** by the change in excitation spectra. The fluorescence from extracellularly facing **FFP-18** was quenched by the membrane impermeant ion Na^{2+} . Ratio fluorescence measurement of **FFP-18** under these conditions permitted the **detection** of **near membrane** Ca^{2+} changes resulting from the release of Ca^{2+} from intracellular stores. **Near membrane** and cytosolic Ca^{2+} changes were **measured** under conditions in which store release and Ca^{2+} influx were triggered by fMLP, thapsigargin or immune complexes. There were significant differences in the timing and magnitude of Ca^{2+} storage site deep within the neutrophil released by thapsigargin and fMLP, but Ca^{2+} near the inner face of the plasma membrane thus provides evidence for the existence of two distinct Ca^{2+} storage locations in neutrophils.

L13 ANSWER 74 OF 113 CA COPYRIGHT 2008 ACS on STN

AN 125:81153 CA

TI Synthesis and characterization of leakage resistant and **near membrane**
 fluorescent calcium **indicator** dyes
 AU Vorndran, Charles
 CS Univ. of Texas, Austin, TX, USA
 SO (1995) 184 pp. Avail.: Univ. Microfilms Int., Order No. DA9617367 From:
 Diss. Abstr. Int., B 1996, 57(1), 62
 TI Synthesis and characterization of leakage resistant and **near membrane**
 fluorescent calcium **indicator** dyes

L13 ANSWER 76 OF 113 CA COPYRIGHT 2008 ACS on STN
 AN 124:4271 CA
 TI New fluorescent calcium **indicators** designed for cytosolic retention or
measuring calcium near membranes
 AU Vorndran, Charles; Minta, Akwasi; Poenie, Martin
 CS Dep. Zool., Univ. Texas, Austin, TX, 78712-1064, USA
 SO Biophysical Journal (1995), 69(5), 2112-24
 AB A new family of fluorescent calcium indicators has been developed based
 on a new analog of BAPTA called FF6. This new BAPTA analog serves as a
 versatile synthetic intermediate for developing Ca²⁺ indicators targeted
 to specific intracellular environments. Two of these new Ca²⁺
 indicators, fura-PE3 and fura-**FFP18**, are described in this report.
 Fura-PE3 is a **zwitterionic** indicator that resists the rapid leakage and
 compartmentalization seen with fura-2 and other polycarboxylate calcium
 indicators. In contrast to results obtained with fura-2, cells loaded
 with PE3 remain brightly loaded and responsive to changes in concn. of
 cytosolic free calcium for hours. Fura-**FFP18** is an amphipathic
 indicator that binds to liposomes and to cell membranes. Studies to
 be detailed later indicate that **FFP18** functions as a **near-membrane** Ca²⁺
indicator and that calcium levels **near** the plasma **membrane** rise faster
 and higher than in the cytosol.

L13 ANSWER 82 OF 113 CA COPYRIGHT 2008 ACS on STN
 AN 120:293334 CA
 TI **Detection** of changes in **near-membrane Ca²⁺** concentration using a novel
 membrane-associated Ca²⁺ indicator
 AU Etter, Elaine F.; Kuhn, Michael A.; Fay, Fredric S.
 CS Med. Sch., Univ. Massachusetts, Worcester, MA, 01605, USA
 SO Journal of Biological Chemistry (1994), 269(13), 10141-9
 AB A Ca²⁺ indicator has been synthesized and characterized which can be
 used to **monitor** rapid changes in the free **Ca²⁺** concn. ([Ca²⁺])
 immediately **adjacent** to cell **membranes**. This indicator, referred to as
 C18-Fura-2, consists of a Fura-2 mol. conjugated to a lipophilic alkyl
 chain which will insert into cell membranes. When assocd. with cell
 membranes in low concns., C18-Fura-2 exhibits an excitation spectrum
 with a large Stokes shift and a single isosbestic point, thus [Ca²⁺] can
 be calcd. ratiometrically. The apparent Ca²⁺ dissocn. const. of cell-
 assocd. C18-Fura-2 is around 150 nM. C18-Fura-2 orients in the cell
 membrane so that the fluorophore is facing the side to which it was
 applied. C18-Fura-2 was used to record rapid changes in intracellular
 [Ca²⁺] which occurred in response to membrane depolarization in isolated
 smooth muscle cells. The initial rise of the [Ca²⁺] transient reported
 by C18-Fura-2 was four to six times faster than the rise of the [Ca²⁺]
 transient reported by cytosolic Fura-2. This result suggests that C18-

Fura-2 was located at the plasma membrane near sites of Ca²⁺ influx and indicates that membrane-assocd. Ca²⁺ indicators can be used to detect rapid, localized changes in [Ca²⁺] which are obscured in signals recorded using water-sol., bulk cytosolic fluorescent Ca²⁺ indicators.

=> log y

STN INTERNATIONAL LOGOFF AT 10:18:45 ON 01 JAN 2008

=> d his

(FILE 'HOME' ENTERED AT 06:10:52 ON 01 JAN 2008)

FILE 'REGISTRY' ENTERED AT 06:11:10 ON 01 JAN 2008

L1 STRUCTURE UPLOADED

L2 STRUCTURE UPLOADED

L3 0 S L1

L4 1 S L2

L5 20 S L2 FULL

FILE 'CA' ENTERED AT 06:18:38 ON 01 JAN 2008

L6 336 S L5

L7 2340 S FLUO

L8 74 S L7 AND DERIVATI?

L9 301 S L6 AND(CALCIUM OR CA OR CA2)

L10 8 S L9 AND LEAK?

L11 80 S L9 AND MEMBRANE

L12 27 S L6 AND DERIVATI?

L13 93 S L6-7 AND MODIF?

L14 240 S L8,L10-13

L15 165 S L14 AND PY<2004

L16 40 S L14 AND PY<2006 AND PATENT/DT

L17 3 S L6 AND MINTA ?/AU

L18 10 S L6-7(5A) (DERIVATI? OR MODIF?)

L19 50 S L6-7(8A) (LEAK? OR MEMBRANE)

L20 49 S L18-19 AND PY<2005

FILE 'BIOSIS' ENTERED AT 06:36:43 ON 01 JAN 2008

L21 50 S L20

FILE 'MEDLINE' ENTERED AT 06:37:01 ON 01 JAN 2008

L22 38 S L20

FILE 'CA, BIOSIS, MEDLINE' ENTERED AT 06:38:45 ON 01 JAN 2008

L23 216 DUP REM L15 L16 L17 L20 L21 L22 (129 DUPLICATES REMOVED)

=> d bib,ab,kwic 1-216 123

L23 ANSWER 16 OF 216 BIOSIS on STN

AN 2004:288730 BIOSIS

TI Near-Membrane Ca²⁺ Measurement with Novel Fluorochromes in Arterial Myocytes.

AU Cavalli, Maurizio [Reprint Author]; Lee, Moo Yeol; Ohkura, Masamichi; Song, Hong; Zhang, Jin; Kinsey, Stephen P; Nakai, Junichi; Kotlikoff, Michael I; Blaustein, Mordecai P

CS Physiol, U Maryland Med Sch, 655 W. Baltimore St, Baltimore, MD, 21201, USA mcava001@umaryland.edu

SO FASEB Journal, (2004) Vol. 18, No. 4-5, pp. Abst. 829.11.

<http://www.fasebj.org/>. e-file. Meeting Info.: FASEB Meeting on Experimental Biology: Translating the Genome. Washington, District of

Columbia, USA. April 17-21, 2004. FASEB.

AB PLasmERosomes, Ca²⁺ signaling complexes, consist of certain plasma membrane (PM) microdomains, the subjacent "junctional" sarco- (or endo-) plasmic reticulum, and the intervening cytosol. Ca²⁺ concentrations in these tiny sub-PM cytosolic spaces ((Ca²⁺)SPM) are apparently regulated independently of the Ca²⁺ in bulk cytosol. Novel "near-membrane" Ca²⁺ indicators should enable us to measure (Ca²⁺)SPM and thereby study PLasmERosome function directly. Fluo-MOMO-AM (TefLabs, Austin, TX), a fluorochrome based on Fluo-4-AM, was loaded into intact rodent small mesenteric arteries (RSMA). Confocal microscopy verified that Fluo-MOMO is anchored to PM and organelle membranes by a hydrophobic tail, and that it detects cytosolic Ca²⁺ signals. We also generated PM-targeted derivatives of G-CaMP, a Ca²⁺-sensitive dye based on green fluorescent protein (Nakai et al., Nature Biotech. 19:137, 2001). We fused the gene for an improved G-CaMP (G-CaMP2; with increased quantum efficiency and extinction coefficient) to the C-terminus of the gene for the Na⁺ pump (1 subunit that is uniformly distributed in the PM. Plasmids were transfected into intact RSMA and primary cultured artery myocytes. Confocal and wide field imaging verified the PM localization of expressed protein and its ability to detect Ca²⁺ signals. A gene for G-CaMP2 fused to the Na/Ca exchanger isoform 1 that is confined to PLasmERosomes was also constructed.

L23 ANSWER 200 OF 216 CA COPYRIGHT 2008 ACS on STN

AN 112:135620 CA

TI Preparation and properties of calcium-specific, long-wavelength indicator dyes

IN Tsien, Roger Yonchien; Minta, Akwasi

PA University of California, Berkeley, USA

SO Eur. Pat. Appl., 27 pp.

PI EP 314480 A2 19890503 EP 1988-310120 19881027
<--

US 5049673 A 19910917 US 1987-115921 19871030
<--

PRAI US 1987-115921 A 19871030

OS MARPAT 112:135620

AB The title dyes I and II [E1, E2 = H, Me, Et, CH₂OH, CO₂H, CH₂CO₂H, or E1E2 = (CH₂)mVCH₂)n (sic; m, n = 1, 2; V = CH₂, O, NH, NMe, S, SS); W = H, OH, CO₂H; X = H, Me, CO₂H, F, Cl, Br, I, NO₂; Y = O, NMe, S, CH₂, CMe₂, CF₂, C:O, bond; Z1, Z2, Z3, Z4 = H, F, Cl, Br, I, Me; Q1, Q2 = R1R2N, R1R2N:+, (R1, R2 = H, Me, Et), OH-, O2-, etc.] and their pharmaceutically acceptable nontoxic salts and esters are provided. Binding of Ca²⁺ increases the fluorescence of the above dyes by up to 40-fold. The Ca²⁺ dissocn. consts. are in the range 0.37-2.3 .mu.M, so that the indicators give better resoln. of high Ca²⁺ concns. than were previously obtainable with predecessor compds. The visible excitation wavelengths of I and II are more convenient for fluorescent microscopy and flow cytometry than the UV required by previous indicators. Thus, III was prepd. from reaction of 2,7-dichloro-3,6-dihydroxyxanth-9-one and an organolithium deriv. (prepn. given) of 1-(2-aminophenoxy)-2-(2-amino-5-phenoxy)ethane, followed by removal of tert-Bu groups. After purifn., extinction coeffs. were 7.9 .times. 10⁴ and 8.3 .times. 10⁴ M⁻¹ cm⁻¹ at 503 and 506 nm, resp., for free and Ca-bound III. Excitation and emission max. for III in the presence of excess Ca were 506 and 526

nm, resp.; quantum efficiencies in the absence of Ca and in the presence of excess Ca were 0.0051 and 0.183, resp. The fluorescence ratio of III in excess Ca vs. no Ca was 36-40. The effective dissociation constant for Ca^{2+} was 0.45 μM .

L23 ANSWER 201 OF 216 CA COPYRIGHT 2008 ACS on STN DUPLICATE 73
AN 111:53566 CA <<LOGINID::20080101>>
TI Fluorescent indicators for cytosolic calcium based on rhodamine and fluorescein chromophores
AU Minta, Akwasi; Kao, Joseph P. Y.; Tsien, Roger Y.
CS Dep. Physiol.-Anat., Univ. California, Berkeley, CA, 94720, USA
SO Journal of Biological Chemistry (1989), 264(14), 8171-8
AB A new group of fluorescent indicators with visible excitation and emission wavelengths was synthesized for measurements of cytosolic free Ca^{2+} . The 5 compounds, rhod-1, rhod-2, fluo-1, fluo-2, and fluo-3, combine the 8-coordinate tetracarboxylate chelating site of 1,2-bis(2-amino-phenoxyethane-N,N,N',N'-tetraacetic acid with a xanthene chromophore to give a rhodamine-like or fluorescein-like fluorophore. Binding of Ca^{2+} increases the fluorescence by up to 40-fold. The Ca^{2+} dissociation constants are in the range 0.37-2.3 μM so that the new indicators should give better resolution of high $[\text{Ca}^{2+}]$ levels than previously obtainable with quin-2 or fura-2. The visible excitation wavelengths of the new compounds are more convenient for fluorescence microscopy and flow cytometry than the UV required by previous indicators. However, the increase in fluorescence of the new dye upon binding Ca is not accompanied by a wavelength shift, so they are unsuitable for measurements using ratios at 2 wavelengths. The most promising dye of this series is fluo-3, which was tested in fibroblasts.

=> log y

STN INTERNATIONAL LOGOFF AT 06:41:47 ON 01 JAN 2008